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- (c) contacting the cell lysate with a separation matrix under conditions suitable for the protein to associate with the separation matrix without diluting the protein prior to the contacting;

- (d) washing the separation matrix; and

- (e) eluting the protein from the separation matrix.

**10.** A method of purifying a protein expressed in a non-native limited solubility form in *E. coli* comprising:

- (a) expressing a protein in a non-native limited solubility form in *E. coli*;

- (b) lysing the *E. coli*;

- (c) solubilizing the expressed protein in a solubilization solution comprising one or more of the following:

- (i) a denaturant;

- (ii) a reductant; and

- (iii) a surfactant;

- (d) forming a refold solution comprising the solubilized protein and a refold buffer, the refold buffer comprising one or more of the following:

- (i) a denaturant;

- (ii) an aggregation suppressor;

- (iii) a protein stabilizer; and

- (iv) a redox component;

- (e) applying the refold solution to a separation matrix under conditions suitable for the protein to associate with the separation matrix and obtaining a purified protein.

**11.** The method of claim **8** or **10**, wherein the non-native limited solubility form is a component of an inclusion body.

**12.** The method of claim **8** or **10**, wherein the protein is a complex protein.

**13.** The method of claim **12**, wherein the complex protein is selected from the group consisting of: a multimeric protein, an antibody, a peptibody, and an Fc fusion protein.

**14.** The method of claim **8** or **10**, wherein the non-native limited solubility form is a component of an inclusion body.

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**15.** The method of claim **8** or **10**, wherein the denaturant in the solubilization solution comprises one or more of: urea, guanidinium salts, dimethyl urea, methylurea, and ethylurea.

**16.** The method of claim **8** or **10**, wherein the reductant comprises one or more of: cysteine, dithiothreitol (DTT), beta-mercaptoethanol, and glutathione.

**17.** The method of claim **8** or **10**, wherein the surfactant comprises one or more of: sarcosyl and sodium dodecylsulfate.

**18.** The method of claim **8** or **10**, wherein the aggregation suppressor is selected from the group consisting of: arginine, proline, polyethylene glycols, nonionic surfactants, ionic surfactants, polyhydric alcohols, glycerol, sucrose, sorbitol, glucose, Tris, sodium sulfate, potassium sulfate, and osmolytes.

**19.** The method of claim **8** or **10**, wherein the protein stabilizer comprises one or more of: arginine, proline, polyethylene glycols, non-ionic surfactants, ionic surfactants, polyhydric alcohols, glycerol, sucrose, sorbitol, glucose, tris, sodium sulfate, potassium sulfate, and osmolytes.

**20.** The method of claim **8** or **10**, wherein the redox component comprises one or more of: glutathione-reduced, glutathione-oxidized, cysteine, cystine, cysteamine, cystamine, and beta-mercaptoethanol.

**21.** The method of claim **8** or **10**, wherein the separation matrix is:

an affinity resin selected from the group consisting of: Protein A, Protein G, and synthetic mimetic affinity resin.

**22.** The method of claim **8** or **10**, wherein the separation matrix is: a non-affinity resin selected from the group consisting of: ion exchange, mixed mode, and a hydrophobic interaction resin.

**23.** The method of claim **8** or **10**, wherein the refold solution is directly applied to the separation matrix.

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